

Applicant : Cawthorne
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COMPLETE LISTING OF ALL CLAIMS, WITH MARKINGS AND STATUS IDENTIFIERS
(Currently amended claims showing deletions by strikethrough and additions by underlining)

1-2 (canceled)

3 (currently amended): A method of ~~claim 2~~ decreasing body weight in a patient, wherein said method comprising administering a therapeutically effective amount of a somatostatin agonist ~~is a somatostatin type-2 receptor agonist to said patient~~.

4 (canceled)

5 (currently amended): A The method of claim 3, wherein said somatostatin type-2 receptor agonist has a Ki of less than 2 nM for the somatostatin type-2 receptor.

6 (canceled)

7 (currently amended): A The method of claim 2 3, wherein said somatostatin agonist is a somatostatin type-2 receptor selective agonist.

8 (canceled)

9 (currently amended): A The method of claim 7, wherein said somatostatin type-2 receptor selective agonist has a Ki for the somatostatin type-2 receptor that is at least 10 times less than the Ki for the somatostatin type-1, type-3, type-4, and type-5 receptors.

10-13 (canceled)

14 (currently amended): A The method of claim 3, wherein said patient is a non-insulin-dependent diabetic human.

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15 (canceled)

16 (currently amended): A The method of claim 5,
wherein said patient is a non-insulin-dependent diabetic human.

17 (canceled)

18 (currently amended): A The method of claim 7,
wherein said patient is a non-insulin-dependent diabetic human.

19 (canceled)

20 (currently amended): A The method of claim 9,
wherein said patient is a non-insulin-dependent diabetic human.

21-22 (canceled)

23 (currently amended): A The method according to
claim 1 3 wherein the somatostatin agonist is

H-D- β -Nal-Cys-Tyr-D-Trp-Lys-Thr-Cys-Thr-NH₂,

H-D-Phe-Cys-Phe-D-Trp-Lys-Thr-Cys- β -Nal-NH₂,

H-D-Phe-Cys-Tyr-D-Trp-Lys-Thr-Cys- β -Nal-NH₂,

H-D- β -Nal-Cys-Phe-D-Trp-Lys-Thr-Cys-Thr-NH₂,

H-D-Phe-Cys-Tyr-D-Trp-Lys-Thr-Pen-Thr-NH₂,

H-D-Phe-Cys-Phe-D-Trp-Lys-Thr-Pen-Thr-NH₂,

H-D-Phe-Cys-Tyr-D-Trp-Lys-Thr-Pen-Thr-OH,

H-D-Phe-Cys-Phe-D-Trp-Lys-Thr-Pen-Thr-OH,

H-Gly-Pen-Phe-D-Trp-Lys-Thr-Cys-Thr-OH,

H-Phe-Pen-Tyr-D-Trp-Lys-Thr-Cys-Thr-OH,

H-Phe-Pen-Phe-D-Trp-Lys-Thr-Pen-Thr-OH,

H-D-Phe-Cys-Phe-D-Trp-Lys-Thr-Cys-Thr-ol

H-D-Phe-Cys-Phe-D-Trp-Lys-Thr-Cys-Thr-NH₂,

H-D-Trp-Cys-Tyr-D-Trp-Lys-Val-Cys-Thr-NH₂,

H-D-Trp-Cys-Phe-D-Trp-Lys-Thr-Cys-Thr-NH₂,

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H-D-Phe-Cys-Tyr-D-Trp-Lys-Val-Cys-Thr-NH₂,
H-D-Phe-Cys-Tyr-D-Trp-Lys-Val-Cys-Trp-NH₂,
H-D-Phe-Cys-Tyr-D-Trp-Lys-Val-Cys-Thr-NH₂,
Ac-D-Phe-Lys*-Tyr-D-Trp-Lys-Val-Asp-Thr-NH₂ (an amide bridge formed between Lys* and Asp),
Ac-hArg(Et)₂-Gly-Cys-Phe-D-Trp-Lys-Thr-Cys-Thr-NH₂,
Ac-D-hArg(Et)₂-Gly-Cys-Phe-D-Trp-Lys-Thr-Cys-Thr-NH₂,
Ac-D-hArg(Bu)-Gly-Cys-Phe-D-Trp-Lys-Thr-Cys-Thr-NH₂,
Ac-D-hArg(Et)₂-Cys-Phe-D-Trp-Lys-Thr-Cys-Thr-NH₂,
Ac-L-hArg(Et)₂-Cys-Phe-D-Trp-Lys-Thr-Cys-Thr-NH₂,
Ac-D-hArg(CH₂CF₃)₂-Cys-Phe-D-Trp-Lys-Thr-Cys-Thr-NH₂,
Ac-D-hArg(CH₂CF₃)₂-Gly-Cys-Phe-D-Trp-Lys-Thr-Cys-Thr-NH₂,
Ac-D-hArg(CH₂CF₃)₂-Gly-Cys-Phe-D-Trp-Lys-Thr-Cys-Phe-NH₂,
Ac-D-hArg(CH₂CF₃)₂-Gly-Cys-Phe-D-Trp-Lys-Thr-Cys-Thr-NHET,
Ac-L-hArg(CH₂CF₃)₂-Gly-Cys-Phe-D-Trp-Lys-Thr-Cys-Thr-NH₂,
Ac-D-hArg(CH₂CF₃)₂-Gly-Cys-Phe-D-Trp-Lys(Me)-Thr-Cys-Thr-NH₂,
Ac-D-hArg(CH₂CF₃)₂-Gly-Cys-Phe-D-Trp-Lys(Me)-Thr-Cys-Thr-NHET,
Ac-hArg(CH₃, hexyl)-Gly-Cys-Phe-D-Trp-Lys-Thr-Cys-Thr-NH₂,
H-hArg(hexyl₂)-Gly-Cys-Phe-D-Trp-Lys-Thr-Cys-Thr-NH₂,
Ac-D-hArg(Et)₂-Gly-Cys-Phe-D-Trp-Lys-Thr-Cys-Thr-NHET,
Ac-D-hArg(Et)₂-Gly-Cys-Phe-D-Trp-Lys-Thr-Cys-Phe-NH₂,
Propionyl-D-hArg(Et)₂-Gly-Cys-Phe-D-Trp-Lys(iPr)-Thr-Cys-Thr-NH₂,
Ac-D-β-Nal-Gly-Cys-Phe-D-Trp-Lys-Thr-Cys-Gly-hArg(Et)₂-NH₂,
Ac-D-Lys(iPr)-Gly-Cys-Phe-D-Trp-Lys-Thr-Cys-Thr-NH₂,
Ac-D-hArg(CH₂CF₃)₂-D-hArg(CH₂CF₃)₂-Gly-Cys-Phe-D-Trp-Lys-Thr-Cys-Thr-NH₂,
Ac-D-hArg(CH₂CF₃)₂-D-hArg(CH₂CF₃)₂-Gly-Cys-Phe-D-Trp-Lys-Thr-Cys-Phe-NH₂,
Ac-D-hArg(Et)₂-D-hArg(Et)₂-Gly-Cys-Phe-D-Trp-Lys-Thr-Cys-Thr-NH₂,
Ac-Cys-Lys-Asn-4-Cl-Phe-Phe-D-Trp-Lys-Thr-Phe-Thr-Ser-D-Cys-NH₂,
H-Bmp-Tyr-D-Trp-Lys-Val-Cys-Thr-NH₂,
H-Bmp-Tyr-D-Trp-Lys-Val-Cys-Phe-NH₂,
H-Bmp-Tyr-D-Trp-Lys-Val-Cys-p-Cl-Phe-NH₂,
H-Bmp-Tyr-D-Trp-Lys-Val-Cys-β-Nal-NH₂,
H-D-β-Nal-Cys-Tyr-D-Trp-Lys-Val-Cys-Thr-NH₂,

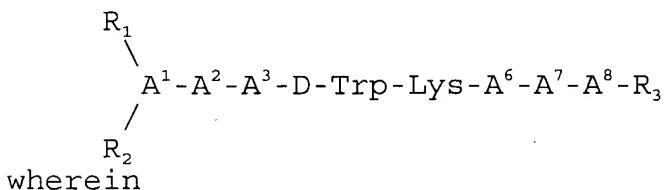
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H-D-Phe-Cys-Tyr-D-Trp-Lys-Abu-Cys-Thr-NH₂,
H-D-Phe-Cys-Tyr-D-Trp-Lys-Abu-Cys-β-Nal-NH₂,
H-pentafluoro-D-Phe-Cys-Tyr-D-Trp-Lys-Val-Cys-Thr-NH₂,
Ac-D-β-Nal-Cys-pentafluoro-Phe-D-Trp-Lys-Val-Cys-Thr-NH₂,
H-D-β-Nal-Cys-Tyr-D-Trp-Lys-Val-Cys-β-Nal-NH₂,
H-D-Phe-Cys-Tyr-D-Trp-Lys-Val-Cys-β-Nal-NH₂,
H-D-β-Nal-Cys-Tyr-D-Trp-Lys-Abu-Cys-Thr-NH₂,
H-D-p-Cl-Phe-Cys-Tyr-D-Trp-Lys-Abu-Cys-Thr-NH₂,
Ac-D-p-Cl-Phe-Cys-Tyr-D-Trp-Lys-Abu-Cys-Thr-NH₂,
H-D-Phe-Cys-β-Nal-D-Trp-Lys-Val-Cys-Thr-NH₂,
H-D-Phe-Cys-Tyr-D-Trp-Lys-Cys-Thr-NH₂,
cyclo(Pro-Phe-D-Trp-N-Me-Lys-Thr-Phe),
cyclo(Pro-Phe-D-Trp-N-Me-Lys-Thr-Phe),
cyclo(Pro-Phe-D-Trp-Lys-Thr-N-Me-Phe),
cyclo(N-Me-Ala-Tyr-D-Trp-Lys-Thr-Phe),
cyclo(Pro-Tyr-D-Trp-Lys-Thr-Phe),
cyclo(Pro-Phe-D-Trp-Lys-Thr-Phe),
cyclo(Pro-Phe-L-Trp-Lys-Thr-Phe) (SEQ ID NO:1),
cyclo(Pro-Phe-D-Trp(F)-Lys-Thr-Phe),
cyclo(Pro-Phe-Trp(F)-Lys-Thr-Phe) (SEQ ID NO:2),
cyclo(Pro-Phe-D-Trp-Lys-Ser-Phe),
cyclo(Pro-Phe-D-Trp-Lys-Thr-p-Cl-Phe),
cyclo(D-Ala-N-Me-D-Phe-D-Thr-D-Lys-Trp-D-Phe),
cyclo(D-Ala-N-Me-D-Phe-D-Val-Lys-D-Trp-D-Phe),
cyclo(D-Ala-N-Me-D-Phe-D-Thr-Lys-D-Trp-D-Phe),
cyclo(D-Abu-N-Me-D-Phe-D-Val-Lys-D-Trp-D-Tyr),
cyclo(Pro-Tyr-D-Trp-t-4-AchxAla-Thr-Phe),
cyclo(Pro-Phe-D-Trp-t-4-AchxAla-Thr-Phe),
cyclo(N-Me-Ala-Tyr-D-Trp-Lys-Val-Phe),
cyclo(N-Me-Ala-Tyr-D-Trp-t-4-AchxAla-Thr-Phe),
cyclo(Pro-Tyr-D-Trp-4-Amphe-Thr-Phe),
cyclo(Pro-Phe-D-Trp-4-Amphe-Thr-Phe),
cyclo(N-Me-Ala-Tyr-D-Trp-4-Amphe-Thr-Phe),
cyclo(Asn-Phe-Phe-D-Trp-Lys-Thr-Phe-Gaba),
cyclo(Asn-Phe-Phe-D-Trp-Lys-Thr-Phe-Gaba-Gaba),

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cyclo(Asn-Phe-D-Trp-Lys-Thr-Phe),
cyclo(Asn-Phe-Phe-D-Trp-Lys-Thr-Phe-NH(CH₂)₄CO),
cyclo(Asn-Phe-Phe-D-Trp-Lys-Thr-Phe-β-Ala),
cyclo(Asn-Phe-Phe-D-Trp-Lys-Thr-Phe-D-Glu)-OH,
cyclo(Phe-Phe-D-Trp-Lys-Thr-Phe),
cyclo(Phe-Phe-D-Trp-Lys-Thr-Phe-Gly),
cyclo(Phe-Phe-D-Trp-Lys-Thr-Phe-Gaba),
cyclo(Asn-Phe-Phe-D-Trp-Lys-Thr-Phe-Gly),
cyclo(Asn-Phe-Phe-D-Trp(F)-Lys-Thr-Phe-Gaba),
cyclo(Asn-Phe-Phe-D-Trp(NO₂)-Lys-Thr-Phe-Gaba),
cyclo(Asn-Phe-Phe-Trp(Br)-Lys-Thr-Phe-Gaba) (SEQ ID NO:3),
cyclo(Asn-Phe-Phe-D-Trp-Lys-Thr-Phe(I)-Gaba),
cyclo(Asn-Phe-Phe-D-Trp-Lys-Thr-Tyr(But)-Gaba),
cyclo(Bmp-Lys-Asn-Phe-Phe-D-Trp-Lys-Thr-Phe-Thr-Pro-Cys)-OH,
cyclo(Bmp-Lys-Asn-Phe-Phe-D-Trp-Lys-Thr-Phe-Thr-Pro-Cys)-OH,
cyclo(Bmp-Lys-Asn-Phe-Phe-D-Trp-Lys-Thr-Phe-Thr-Tpo-Cys)-OH,
cyclo(Bmp-Lys-Asn-Phe-Phe-D-Trp-Lys-Thr-Phe-Thr-MeLeu-Cys)-OH,
cyclo(Phe-Phe-D-Trp-Lys-Thr-Phe-Phe-Gaba),
cyclo(Phe-Phe-D-Trp-Lys-Thr-Phe-D-Phe-Gaba),
cyclo(Phe-Phe-D-Trp(5F)-Lys-Thr-Phe-Phe-Gaba),
cyclo(Asn-Phe-Phe-D-Trp-Lys(Ac)-Thr-Phe-NH-(CH₂)₃-CO),
cyclo(Lys-Phe-Phe-D-Trp-Lys-Thr-Phe-Gaba),
cyclo(Lys-Phe-Phe-D-Trp-Lys-Thr-Phe-Gaba),
cyclo(Orn-Phe-Phe-D-Trp-Lys-Thr-Phe-Gaba),
H-Cys-Phe-Phe-D-Trp-Lys-Thr-Phe-Cys-NH₂,
H-Cys-Phe-Phe-D-Trp-Lys-Ser-Phe-Cys-NH₂,
H-Cys-Phe-Tyr-D-Trp-Lys-Thr-Phe-Cys-NH₂ or
H-Cys-Phe-Tyr(I)-D-Trp-Lys-Thr-Phe-Cys-NH₂.

24 (currently amended): A The method according to
claim 1 3 wherein the somatostatin agonist is



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A¹ is a D- or L- isomer of Ala, Leu, Ile, Val, Nle, Thr, Ser, β -Nal, β -Pal, Trp, Phe, 2,4-dichloro-Phe, pentafluoro-Phe, p-X-Phe, or o-X-Phe, wherein X is CH₃, Cl, Br, F, OH, OCH₃ or NO₂;

A² is Ala, Leu, Ile, Val, Nle, Phe, β -Nal, pyridyl-Ala, Trp, 2,4-dichloro-Phe, pentafluoro-Phe, o-X-Phe, or p-X-Phe, wherein X is CH₃, Cl, Br, F, OH, OCH₃ or NO₂;

A³ is pyridyl-Ala, Trp, Phe, β -Nal, 2,4-dichloro-Phe, pentafluoro-Phe, o-X-Phe, or p-X-Phe, wherein X is CH₃, Cl, Br, F, OH, OCH₃ or NO₂;

A⁶ is Val, Ala, Leu, Ile, Nle, Thr, Abu, or Ser;

A⁷ is Ala, Leu, Ile, Val, Nle, Phe, β -Nal, pyridyl-Ala, Trp, 2,4-dichloro-Phe, pentafluoro-Phe, o-X-Phe, or p-X-Phe, wherein X is CH₃, Cl, Br, F, OH, OCH₃ or NO₂;

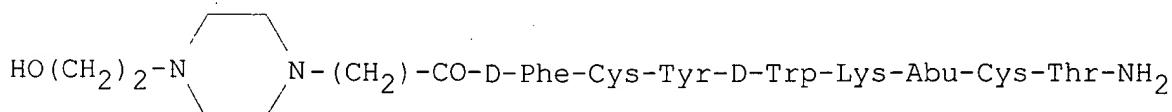
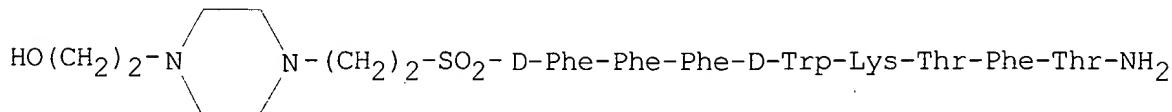
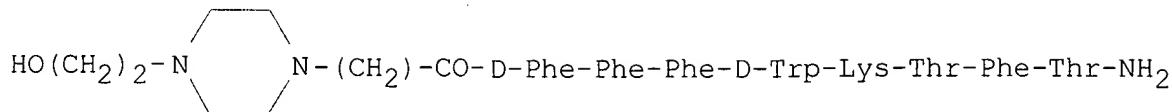
A⁸ is a D- or L-isomer of Ala, Leu, Ile, Val, Nle, Thr, Ser, Phe, β -Nal, pyridyl-Ala, Trp, 2,4-dichloro-Phe, pentafluoro-Phe, p-X-Phe, or o-X-Phe, wherein X is CH₃, Cl, Br, F, OH, OCH₃ or NO₂;

each R₁ and R₂, independently, is H, lower acyl or lower alkyl; and R₃ is OH or NH₂; provided that at least one of A¹ and A⁸ and one of A² and A⁷ must be an aromatic amino acid; and further provided that A¹, A², A⁷ and A⁸ cannot all be aromatic amino acids.

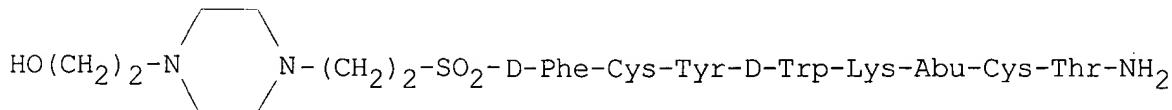
25 (currently amended): A The method according to claim 24 wherein the linear somatostatin agonist is H-D-Phe-p-chloro-Phe-Tyr-D-Trp-Lys-Thr-Phe-Thr-NH₂, H-D-Phe-p-NO₂-Phe-Tyr-D-Trp-Lys-Val-Phe-Thr-NH₂, H-D-Nal-p-chloro-Phe-Tyr-D-Trp-Lys-Val-Phe-Thr-NH₂, H-D-Phe-Phe-Phe-D-Trp-Lys-Thr-Phe-Thr-NH₂ , H-D-Phe-Phe-Tyr-D-Trp-Lys-Val-Phe-Thr-NH₂, H-D-Phe-p-chloro-Phe-Tyr-D-Trp-Lys-Val-Phe-Thr-NH₂ or H-D-Phe-Ala-Tyr-D-Trp-Lys-Val-Ala- β -D-Nal-NH₂.

26 (currently amended): A The method according to claim 13 wherein the somatostatin agonist is

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or



27 (canceled)

28 (currently amended) : A The method according to
claim 3 wherein said patient is obese.

29 (canceled)

30 (currently amended) : A The method according to
claim 7 wherein said patient is obese.

31-32 (canceled)